Response and Amendment under 37 CFR § 1.111 Application No. 09/516,194 Page 2 of 14

II. Amendments to the Claims

This listing of claims will replace all prior versions, and listings, of claims in the application.

Listing of Claims:

- 1. (Canceled)
- 2. (Currently Amended) A compound of formula (I) or a pharmaceutically acceptable salt thereof, wherein the compound of formula (I) is:

$$R_1$$
 R_2
 R_3
 R_4
 R_7
 R_7
 R_8
 R_8
 R_9
 R_9

wherein the dotted lines indicate a single or a double bond;

$$R_1$$
 is $-OD_1$ or $-Cl$;

 R_2 and R_8 are a hydrogen; or R_1 and R_2 taken together are $=CH_2$ or =O;

R₃ and R₄ are each independently a hydrogen, -OD₁ or -CH₃;

R₅ and R₆ are each independently a hydrogen, -OD₁, -CH₃, -OCH₃ or -CH=CH₂;

 R_7 is a hydrogen or $-OD_1$;

 R_9 is hydrogen or absent when the carbon to which it is attached is the central carbon of an allene functionality; or R_8 and R_9 taken together with the chain to which they are attached form a substituted benzene ring with the proviso that R_1 is an oxygen atom which is attached to the carbon atom at the position of the benzene ring defined by B;

A is
$$-CH=$$
, $-CH_2$, $-S-$, or $-O-$;

B is
$$-CH=$$
, $-CH_2$, $-S-$, or $-C(O)-$;

X is
$$-CH_2OR_{11}$$
, $-C(O)OR_{11}$ or $-C(O)N(D_1)R_{12}$;

R₁₁ is D₁, a lower alkyl group, or

Response and Amendment under 37 CFR § 1.111 Application No. 09/516,194 Page 3 of 14

 R_{12} is $-S(O)_2CH_3$ or $-C(O)CH_3$;

Z is (a) an ethyl, (b) a butyl, (c) a hexyl, (d) a benzyl,

R₁₃ is a hydrogen or -Cl;

 D_1 is a hydrogen or D; with the proviso that at least one D_1 in formula (I) must be D;

D is Q or K;

Q is -NO or -NO₂;

K is $-W_a-E_b-(C(R_e)(R_f))_p-E_c-(C(R_e)(R_f))_x-W_d-(C(R_e)(R_f))_y-W_i-E_j-W_g-(C(R_e)(R_f))_z-T-Q;$ with the proviso that when X is $-C(O)OD_1$ and D_1 is K, then K is not an alkyl, branched alkyl or cycloalkyl mononitrate; a benzoic acid substituted benzyloxy mononitrate; the regioisomeric esters of glycerol dinitrate and oligomers thereof;

a, b, c, d, g, i and j are each independently an integer from 0 to 3;

p, x, y and z are each independently an integer from 0 to 10;

W at each occurrence is independently -C(O)-, -C(S)-, -T-, -(C(R_e)(R_f))_h-, an alkyl group, an aryl group, a heterocyclic ring, an arylheterocyclic ring, or -(CH₂CH₂O)_q-;

E at each occurrence is independently -T-, an alkyl group, an aryl group, $-(C(R_e)(R_f))_h$ -, a heterocyclic ring, an arylheterocyclic ring, or $-(CH_2CH_2O)_q$ -;

h is an integer form 1 to 10;

q is an integer from 1 to 5;

Response and Amendment under 37 CFR § 1.111 Application No. 09/516,194 Page 4 of 14

Re and Rf are each independently a hydrogen, an alkyl, a cycloalkoxy, a halogen, a hydroxy, an hydroxyalkyl, an alkoxyalkyl, an arylheterocyclic ring, an alkylaryl, a cycloalkylalkyl, a heterocyclicalkyl, an alkoxy, a haloalkoxy, an amino, an alkylamino, a dialkylamino, an arylamino, a diarylamino, an alkylarylamino, an alkoxyhaloalkyl, a haloalkoxy, a sulfonic acid, a sulfonic ester, an alkylsulfonic acid, an arylsulfonic acid, an arylalkoxy, an alkylthio, a cycloalkylthio, a cycloalkenyl, a cyano, an aminoalkyl, an aminoaryl, an aryl, an arylalkyl, an alkylaryl, a carboxamido, a alkylcarboxamido, an arylcarboxamido, an amidyl, a carboxyl, a carbamoyl, a carbamate, an alkylcarboxylic acid, an arylcarboxylic acid, an arylcarboxylic ester, an alkylcarboxylic ester, an arylcarboxylic ester, a haloalkoxy, a sulfonamido, an alkylsulfonamido, an arylsulfonamido, a sulfonic ester, a urea, a phosphoryl, a nitro, -T-Q, or (C(Re)(Rf))k-T-Q, or Re and Rf taken together with the carbons to which they are attached form a carbonyl, a methanthial, a heterocyclic ring, a cycloalkyl group or a bridged cycloalkyl group;

k is an integer from 1 to 3;

T at each occurrence is independently a covalent bond, a carbonyl, an oxygen, $-S(O)_{o}$ - or $-N(R_a)R_i$ -;

o is an integer from 0 to 2;

R_a is a lone pair of electrons, a hydrogen or an alkyl group;

 R_i is a hydrogen, an alkyl, an aryl, an alkylcarboxylic acid, an arylcarboxylic acid, an alkylcarboxylic ester, an arylcarboxylic ester, an alkylcarboxamido, an arylcarboxamido, an alkylaryl, an alkylsulfinyl, an alkylsulfinyl, an arylsulfinyl, an arylsulfonyl, a sulfonamido, a carboxamido, a carboxylic ester, an amino alkyl, an amino aryl, $-CH_2-C(T-Q)(R_e)(R_f)$, or $-(N_2O_2-)^{-\bullet}M^+$, wherein M^+ is an organic or inorganic cation; with the proviso that when R_i is $-CH_2-C(T-Q)(R_e)(R_f)$ or $-(N_2O_2)^{-\bullet}M^+$, or R_e or R_f are T-Q or $(C(R_e)(R_f))_k-T-Q$, then the "-T-Q" subgroup can be a hydrogen, an alkyl, an alkoxy, an alkoxyalkyl, an aminoalkyl, a hydroxy, a heterocyclic ring or an aryl group;

with the proviso that the compound of formula (I) has at least one NO group or at least one NO₂ group linked through an oxygen atom, a nitrogen atom or a sulfur atom.

3. (Currently Amended) The compound of claim 2, wherein the compound of formula (I) comprising at least one NO group, at least one NO₂ group, or at least one NO and

Response and Amendment under 37 CFR § 1.111 Application No. 09/516,194 Page 5 of 14

NO₂ group is a nitrosated arbaprostil, a nitrosylated arbaprostil, a nitrosated and nitrosylated arbaprostil, a nitrosated alprostadil, a nitrosylated alprostadil, a nitrosated and nitrosylated alprostadil, a nitrosated beraprost, a nitrosylated beraprost, a nitrosated and nitrosylated beraprost, a nitrosated carboprost, a nitrosylated carboprost, a nitrosated and nitrosylated carboprost, a nitrosated cloprostenol, a nitrosylated cloprostenol, a nitrosated and nitrosylated cloprostenol, a nitrosated dimoxaprost, a nitrosylated dimoxaprost, a nitrosated and nitrosylated dimoxaprost, a nitrosated enprostil, a nitrosylated enprostil, a nitrosated and nitrosylated enprostil, a nitrosated enisoprost, a nitrosylated enisoprost, a nitrosated and nitrosylated enisoprost, a nitrosated fluprostenol, a nitrosylated fluprostenol, a nitrosated and nitrosylated fluprostenol, a nitrosated fenprostalene, a nitrosylated fenprostalene, a nitrosated and nitrosylated fenprostalene, a nitrosated gemeprost, a nitrosylated gemeprost, a nitrosated and nitrosylated gemeprost, a nitrosated latanaprost, a nitrosylated latanaprost, a nitrosated and nitrosylated latanaprost, a nitrosated limaprost, a nitrosylated limaprost, a nitrosated and nitrosylated limaprost, a nitrosated meteneprost, a nitrosylated meteneprost, a nitrosated and nitrosylated meteneprost, a nitrosated mexiprostil, a nitrosylated mexiprostil, a nitrosated and nitrosylated mexiprostil, a nitrosated misoprostol, a nitrosylated misoprostol, a nitrosylated misoprostol, a nitrosated misoprost, a nitrosylated misoprost, a nitrosated and nitrosylated misoprost, a nitrosated misoprostol acid, a nitrosylated misoprostol acid, a nitrosated and nitrosylated misoprostol acid, a nitrosated nocloprost, a nitrosylated nocloprost, a nitrosated and nitrosylated nocloprost, a nitrosated ornoprostil, a nitrosylated ornoprostil, a nitrosated and nitrosylated ornoprostil, a nitrosated prostalene, a nitrosylated prostalene, a nitrosated and nitrosylated prostalene, a nitrosated PGE₁, a nitrosylated PGE₁, a nitrosated and nitrosylated PGE₁, a nitrosated PGE₂, a nitrosylated PGE₂, a nitrosated and nitrosylated PGE₂, a nitrosated PGF₁, a nitrosylated PGF₁, a nitrosylated PGF₁, a nitrosylated PGF_{2 α}, a nitrosylated $PGF_{2\alpha}$, a nitrosated and nitrosylated $PGF_{2\alpha}$, a nitrosated rioprostil, a nitrosylated rioprostil, a nitrosated and nitrosylated rioprostil, a nitrosated rosaprostol, a nitrosylated rosaprostol, a nitrosated and nitrosylated rosaprostol, a nitrosated remiprostol, a nitrosylated remiprostol, a nitrosated and nitrosylated remiprostol, a nitrosated sulprostone, a nitrosylated sulprostone, a nitrosated and nitrosylated sulprostone, a nitrosated trimoprostil, a nitrosylated trimoprostil, a nitrosated and nitrosylated trimoprostil, a nitrosated tiprostanide, a nitrosylated tiprostanide, a

nitrosated and nitrosylated tiprostanide, a nitrosated unoprostone, a nitrosylated unoprostone, a nitrosated and nitrosylated unoprostone, a nitrosated viprostol, a nitrosylated viprostol, a nitrosated and nitrosylated viprostol or a mixture thereof.

- 4. (Original) A composition comprising the compound of claim 2 and a pharmaceutically acceptable carrier.
- 5. (Original) A method for treating a sexual dysfunction in a patient in need thereof comprising administering to the patient a therapeutically effective amount of the composition of claim 4.
 - 6. (Original) The method of claim 5, wherein the patient is female.
 - 7. (Original) The method of claim 5, wherein the patient is male.
- 8. (Original) The method of claim 5, wherein the composition is administered orally, by intracavernosal injection, by transurethral application, or by transdermal application.
 - 9. (Cancelled)
- 10. (Original) The composition of claim 4, further comprising at least one vasoactive agent or a pharmaceutically acceptable salt thereof.
- 11. (Original) The composition of claim 10, wherein the vasoactive agent is a potassium channel activator, a calcium channel blocker, an α -blocker, a β -blocker, a phosphodiesterase inhibitor, adenosine, an ergot alkaloid, a vasoactive intestinal peptide, a dopamine agonist, an opioid antagonist, an endothelin antagonist or a mixture thereof.
- 12. (Original) The composition of claim 10, wherein the vasoactive agent is an α -blocker or a phosphodiesterase inhibitor.
- 13. (Original) The composition of claim 12, wherein the α -blocker is phentolamine, prazosin, doxazosin, terazosin, yohimbine or moxisylyte and the phosphodiesterase inhibitor is papaverine, zaprinast, sildenafil or IC 351, or a mixture thereof.
- 14. (Original) A method for treating a sexual dysfunction in a patient in need thereof comprising administering to the patient a therapeutically effective amount of the composition of claim 10.
 - 15. (Original) The method of claim 14, wherein the patient is female.
 - 16. (Original) The method of claim 14, wherein the patient is male.

Response and Amendment under 37 CFR § 1.111 Application No. 09/516,194 Page 7 of 14

- 17. (Original) The method of claim 14, wherein the composition is administered orally, by intracavernosal injection, by transurethral application or by transdermal application.
 - 18. (Canceled)
- 19. (Previously Presented) A composition comprising at least one compound of claim 2 or a pharmaceutically acceptable salt thereof, and at least one compound that donates, transfers or releases nitric oxide, or induces the production of endogenous nitric oxide or endothelium-derived relaxing factor, or is a substrate for nitric oxide synthase.
- 20. (Previously Presented) The composition of claim 19, further comprising a pharmaceutically acceptable carrier.
- 21. (Original) The composition of claim 19, wherein the compound that donates, transfers, or releases nitric oxide, or induces the production of endogenous nitric oxide or endothelium-derived relaxing factor or is a substrate for nitric oxide synthase is an S-nitrosothiol.
- 22. (Original) The composition of claim 21, wherein the S-nitrosothiol is S-nitroso-N-acetylcysteine, S-nitroso-captopril, S-nitroso-N-acetylpenicillamine, S-nitroso-homocysteine, S-nitroso-cysteine or S-nitroso-glutathione.
 - 23. (Original) The composition of claim 21, wherein the S-nitrosothiol is:
 - (i) $HS(C(R_e)(R_f))_mSNO$;
 - (ii) $ONS(C(R_e)(R_f))_mR_e$; and
 - (iii) $H_2N-CH(CO_2H)-(CH_2)_m-C(O)NH-CH(CH_2SNO)-C(O)NH-CH_2-CO_2H$;

wherein m is an integer from 2 to 20; R_e and R_f are each independently a hydrogen, an alkyl, a cycloalkoxy, a halogen, a hydroxy, an hydroxyalkyl, an alkoxyalkyl, an arylheterocyclic ring, an alkylaryl, a cycloalkylalkyl, a heterocyclicalkyl, an alkoxy, a haloalkoxy, an amino, an alkylamino, a dialkylamino, an arylamino, a diarylamino, an alkylarylamino, an alkoxyhaloalkyl, a haloalkoxy, a sulfonic acid, a sulfonic ester, an alkylsulfonic acid, an arylalkoxy, an alkylthio, an arylthio, a cycloalkylthio, a cycloalkenyl, a cyano, an aminoalkyl, an aminoaryl, an aryl, an arylalkyl, an alkylaryl, a carboxamido, a alkylcarboxamido, an arylcarboxamido, an amidyl, a carboxyl, a carbamoyl, a carbamate, an alkylcarboxylic acid, an arylcarboxylic acid, an arylcarboxylic ester, an arylcarboxylic ester, a haloalkoxy, a sulfonamido, an alkylsulfonamido, an arylsulfonamido, a sulfonic ester, a urea, a phosphoryl, a nitro, -T-Q, or $(C(R_e)(R_f))_k$ -T-Q, or

Response and Amendment under 37 CFR § 1.111 Application No. 09/516,194 Page 8 of 14

 R_e and R_f taken together with the carbons to which they are attached form a carbonyl, a methanthial, a heterocyclic ring, a cycloalkyl group or a bridged cycloalkyl group; Q is -NO or -NO₂; and T is independently a covalent bond, a carbonyl, an oxygen, -S(O)_o- or -N(R_a)R_i-, wherein o is an integer from 0 to 2, R_a is a lone pair of electrons, a hydrogen or an alkyl group; R_i is a hydrogen, an alkyl, an aryl, an alkylcarboxylic acid, an aryl carboxylic acid, an alkylcarboxylic ester, an arylcarboxylic ester, an alkylcarboxamido, an arylcarboxamido, an alkylsulfinyl, an alkylsulfinyl, an arylsulfinyl, an arylsulfonyl, a sulfonamido, a carboxamido, a carboxylic ester, an amino alkyl, an amino aryl, -CH₂-C(T-Q)(R_e)(R_f), or -(N_2O_2 -) $^{\bullet}$ M $^+$, wherein M $^+$ is an organic or inorganic cation; with the proviso that when R_i is -CH₂-C(T-Q)(R_e)(R_f) or -(N_2O_2 -) $^{\bullet}$ M $^+$; then "-T-Q" can be a hydrogen, an alkyl group, an alkoxyalkyl group, an aminoalkyl group, a hydroxy group or an aryl group.

- 24. (Original) The composition of claim 19, wherein the compound that donates, transfers, or releases nitric oxide, or induces the production of endogenous nitric oxide or endothelium-derived relaxing factor, or is a substrate for nitric oxide synthase, is L-arginine, L-homoarginine, N-hydroxy-L-arginine, nitrosated L-arginine, nitrosylated L-arginine, nitrosylated L-arginine, nitrosylated N-hydroxy-L-arginine, citrulline, ornithine, glutamine, lysine, polypeptides comprising at least one of these amino acids or inhibitors of the enzyme arginase.
- 25. (Previously Presented) The composition of claim 19, wherein the compound that donates, transfers, or releases nitric oxide, or induces the production of endogenous nitric oxide or endothelium-derived relaxing factor, or is a substrate for nitric oxide synthase is:
 - (i) a compound that comprises at least one ON-O-, ON-N- or ON-C- group;
- (ii) a compound that comprises at least one O_2N -O-, O_2N -N-, O_2N -S- or $-O_2N$ -C- group;
- (iii) a N-oxo-N-nitrosoamine having the formula: R¹R²-N-N(O-M⁺)-NO, wherein R¹ and R² are each independently a polypeptide, an amino acid, a sugar, an oligonucleotide, a straight or branched, saturated or unsaturated, aliphatic or aromatic, substituted or unsubstituted hydrocarbon, or a heterocyclic group, and M⁺ is an organic or inorganic cation.
- 26. (Original) The composition of claim 25, wherein the compound comprising at least one ON-O-, ON-N- or ON-C- group is an ON-O-polypeptide, an ON-N-polypeptide, an

Response and Amendment under 37 CFR § 1.111 Application No. 09/516,194 Page 9 of 14

ON-C-polypeptide, an ON-O-amino acid, an ON-N-amino acid, an ON-C-amino acid, an ON-O-sugar, an ON-N-sugar, an ON-C-sugar, an ON-O-oligonucleotide, an ON-N-oligonucleotide, an ON-C-oligonucleotide, a straight or branched, saturated or unsaturated, substituted or unsubstituted, aliphatic or aromatic ON-O-hydrocarbon, a straight or branched, saturated or unsubstituted, aliphatic or aromatic ON-N-hydrocarbon, a straight or branched, saturated or unsubstituted or unsubstituted, aliphatic or aromatic ON-C-hydrocarbon, an ON-O-heterocyclic compound, an ON-N-heterocyclic compound or a ON-C-heterocyclic compound.

- 27. (Original) The composition of claim 25, wherein compound comprising at least one O₂N-O-, O₂N-N-, O₂N-S- or O₂N-C- group is an O₂N-O-polypeptide, an O₂N-N-polypeptide, an O₂N-S-polypeptide, an O₂N-C-polypeptide, an O₂N-O-amino acid, O₂N-N-amino acid, O₂N-S-amino acid, an O₂N-C-amino acid, an O₂N-O-sugar, an O₂N-N-sugar, O₂N-S-sugar, an O₂N-C-sugar, an O₂N-O-oligonucleotide, an O₂N-N-oligonucleotide, an O₂N-S-oligonucleotide, an O₂N-C-oligonucleotide, a straight or branched, saturated or unsaturated, aliphatic or aromatic, substituted or unsubstituted O₂N-O-hydrocarbon, a straight or branched, saturated or unsaturated, aliphatic or aromatic, substituted or unsubstituted O₂N-N-hydrocarbon, a straight or branched, saturated or unsaturated, aliphatic or aromatic, substituted or unsubstituted or unsubst
- 28. (Original) A method for treating a sexual dysfunction in a patient in need thereof comprising administering to the patient a therapeutically effective amount of the composition of claim 19.
 - 29. (Original) The method of claim 28, wherein the patient is female.
 - 30. (Original) The method of claim 28, wherein the patient is male.
- 31. (Original) The method of claim 28, wherein the composition is administered orally, by intracavernosal injection, by transurethral application or by transdermal application.
 - 32. (Canceled)
- 33. (Original) The composition of claim 19, further comprising at least one vasoactive agent or a pharmaceutically acceptable salt thereof.

Response and Amendment under 37 CFR § 1.111 Application No. 09/516,194 Page 10 of 14

- 34. (Original) The composition of claim 33, wherein the vasoactive agent is a potassium channel activator, a calcium channel blocker, an α -blocker, a β -blocker, a phosphodiesterase inhibitor, adenosine, an ergot alkaloid, a vasoactive intestinal peptide, a dopamine agonist, an opioid antagonist, an endothelin antagonist or a mixture thereof.
- 35. (Original) The composition of claim 34, wherein the vasoactive agent is an α -blocker or a phosphodiesterase inhibitor.
- 36. (Original) The composition of claim 35, wherein the α -blocker is phentolamine, prazosin, doxazosin, terazosin, yohimbine or moxisylyte and the phosphodiesterase inhibitor is papaverine, zaprinast, sildenafil or IC 351, or a mixture thereof.
- 37. (Original) A method for treating a sexual dysfunction in a patient in need thereof comprising administering to the patient a therapeutically effective amount of the composition of claim 33.
 - 38. (Original) The method of claim 37, wherein the patient is female.
 - 39. (Original) The method of claim 37, wherein the patient is male.
- 40. (Original) The method of claim 37, wherein the composition is administered orally, by intracavernosal injection, by transurethral application or by transdermal application.

Claims 41-103 (Cancelled)

- 104. (Original) A kit comprising at least one compound of claim 2 and at least one compound that donates, transfers or releases nitric oxide, or induces the production of endogenous nitric oxide or endothelium-derived relaxing factor, or is a substrate for nitric oxide synthase.
- 105. (Original) The kit of claim 104, wherein the compound of claim 2 and the at least one compound that donates, transfers or releases nitric oxide, induces the production of endogenous nitric oxide or endothelium-derived relaxing factor, or is a substrate for nitric oxide synthase are separate components in the kit or are in the form of a composition in the kit.
 - 106. (Original) The kit of claim 104, further comprising at least one vasoactive agent.
- 107. (Currently Amended) A prostaglandin comprising at least one NO group or a pharmaceutically acceptable salt thereof; wherein the at least one NO group is linked to the prostaglandin compound through an oxygen atom, a nitrogen atom or a sulfur atom.